

Substitute for form 1449A/PTO  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  <i>(Use as many sheets as necessary)</i>				<b>Complete if Known</b>	
				Application Number	10/678,765
				Filing Date	October 2, 2003
				First Named Inventor	Serbedzija, George N.
				Art Unit	1632
Examiner Name	Valarie E. Bertoglio				
Attorney Docket Number	018852-000511US				
Sheet	1	of	5		

U.S. PATENT DOCUMENTS					
Examiner Initials*	Cite No. <sup>1</sup>	Document Number	Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear
		Number Kind Code <sup>2</sup> (if known)			

FOREIGN PATENT DOCUMENTS								
Examiner Initials*	Cite No. <sup>1</sup>	Foreign Patent Document			Publication Date MM-DD-YYYY	Name of Patentee or Applicant of Cited Document	Pages, Columns, Lines, Where Relevant Passages or Relevant Figures Appear	T <sup>6</sup>
		Country Code <sup>3</sup>	Number <sup>4</sup>	Kind Code <sup>5</sup> (if known)				
	230	EP	1 644 733	B1	04-11-2007	Daniolabs Limited		<input type="checkbox"/>

Examiner Signature		Date Considered	
-----------------------	--	--------------------	--

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant. <sup>1</sup> Applicant's unique citation designation number (optional). <sup>2</sup> Kind Codes of U.S. Patent Documents at [www.uspto.gov](http://www.uspto.gov) or MPEP 901.04. <sup>3</sup> Enter Office that issued the document, by the two-letter code (WIPO Standard ST.3). <sup>4</sup> For Japanese patent documents, the indication of the year of the reign of the Emperor must precede the serial number of the patent document. <sup>5</sup> Kind of document by the appropriate symbols as indicated on the document under WIPO Standard ST. 16 if possible. <sup>6</sup> Applicant is to place a check mark here if English language Translation is attached.

Substitute for form 1449B/PTO  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (Use as many sheets as necessary)			<b>Complete if Known</b>		
			Application Number	10/678,765	
			Filing Date	October 2, 2003	
			First Named Inventor	Serbedzija, George N.	
			Art Unit	1632	
			Examiner Name	Valarie E. Bertoglio	
Sheet	2	of	5	Attorney Docket Number	018852-000511US

NON PATENT LITERATURE DOCUMENTS				
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>	
	202	ALEXANDRE et al., "Somatotopy of the lateral line projection in larval zebrafish," <u>PNAS</u> , 96:7558-7562 (1999).	<input type="checkbox"/>	
	194	ANICHTCHIK et al., "Neurochemical and behavioural changes in zebrafish <i>Danio rerio</i> after systemic administration of 6-hydroxydopamine and 1-methyl-4-phenyl-1,2,3,6-tetrahydropyridine," <u>Journal of Neurochemistry</u> , 88:443-453 (2004).	<input type="checkbox"/>	
	208	ANKLEY et al., "Small Fish Models for Identifying and Assessing the Effects of Endocrine-disrupting Chemicals," <u>ILAR Journal</u> , 45(4):469-483 (2004).	<input type="checkbox"/>	
	209	BEHRA et al., "The Use of Zebrafish Mutants to Identify Secondary Target Effects of Acetylcholine Esterase Inhibitors," <u>Toxicological Sciences</u> , 77:325-333 (2004).	<input type="checkbox"/>	
	210	BERNSTEIN et al., "The Blood-Brain Barrier of Fish," <u>Experimental Neurology</u> , 11:464-473 (1965).	<input type="checkbox"/>	
	211	BRETAUD et al., "Sensitivity of zebrafish to environmental toxins implicated in Parkinson's disease," <u>Neurotoxicology and Teratology</u> , 26:857-864 (2004).	<input type="checkbox"/>	
	212	BRUSTEIN et al., "In vivo monitoring of neuronal network activity in zebrafish by two-photon Ca <sup>2+</sup> imaging," <u>Eur J Physiol</u> , 446:766-773 (2003).	<input type="checkbox"/>	
	213	CHANG et al., "Developmental Toxicity of Arecoline, the Major Alkaloid in Betel Nuts, in Zebrafish Embryos," <u>Birth Defects Research</u> , (Part A) 70:28-36 (2004).	<input type="checkbox"/>	
	191	CSERR et al., "Blood-brain interfaces in vertebrates: a comparative approach," <u>Am J Physiol</u> , 246:R277-R288 (1984).	<input type="checkbox"/>	
	214	DARLAND et al., "Behavioral screening for cocaine sensitivity in mutagenized zebrafish," <u>PNAS</u> , 98(20):11691-11696 (2001).	<input type="checkbox"/>	
	215	DEN HERTOOG, Jeroen, "Chemical Genetics: Drug Screens in Zebrafish," <u>Bioscience Reports</u> , 25(5/6):289-297 (2005).	<input type="checkbox"/>	
Examiner Signature			Date Considered	

\* EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup> Applicant's unique citation designation number (optional). <sup>2</sup> Applicant is to place a check mark here if English language Translation is attached.

Substitute for form 1449B/PTO  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (Use as many sheets as necessary)			<b>Complete if Known</b>		
			Application Number	10/678,765	
			Filing Date	October 2, 2003	
			First Named Inventor	Serbedzija, George N.	
			Art Unit	1632	
			Examiner Name	Valarie E. Bertoglio	
Sheet	3	of	5	Attorney Docket Number	018852-000511US

NON PATENT LITERATURE DOCUMENTS				
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>	
	200	DONG et al., "Role of Aryl Hydrocarbon Receptor in Mesencephalic Circulation Failure and Apoptosis in Zebrafish Embryos Exposed to 2,3,7,8-Tetrachlorodibenzo-p-Dioxin," <u>Toxicological Sciences</u> , 77:109-116 (2004).	<input type="checkbox"/>	
	216	DONG et al., "2,3,7,8-Tetrachlorodibenzo-p-dioxin Toxicity in the Zebrafish Embryo: Local Circulation Failure in the Dorsal midbrain Is Associated with Increased Apoptosis," <u>Toxicological Sciences</u> , 69:191-201 (2002).	<input type="checkbox"/>	
	217	FETCHO et al., "Visualization of Active Neural Circuitry in the Spinal Cord of Intact Zebrafish," <u>Journal of Neurophysiology</u> , 73(1):399-406 (1995).	<input type="checkbox"/>	
	206	FRIEDLAND et al., "Development of an Anti-A $\beta$ Monoclonal Antibody for In Vivo Imaging of Amyloid Angiopathy in Alzheimer's Disease," <u>Molecular Neurobiology</u> , 9:107-113 (1994).	<input type="checkbox"/>	
	218	GELING et al., "A $\gamma$ -secretase inhibitor blocks Notch signaling <i>in vivo</i> and causes a severe neurogenic phenotype in zebrafish," <u>EMBO Reports</u> , 3(7):688-694 (2002).	<input type="checkbox"/>	
	219	GERLAI et al., "Drinks like a fish: zebra fish ( <i>Danio rerio</i> ) as a behavior genetic model to study alcohol effects," <u>Pharmacology, Biochemistry, and Behavior</u> , 67:773-782 (2000).	<input type="checkbox"/>	
	197	GUPTA et al., "Neuroprotective Role of Melatonin in Oxidative Stress Vulnerable Brain," <u>Indian J. Physiol Pharmacol</u> , 47(4):373-386 (2003).	<input type="checkbox"/>	
	220	HILL et al., "Neurodevelopmental Defects in Zebrafish ( <i>Danio rerio</i> ) at Environmentally Relevant Dioxin (TCDD) Concentrations," <u>Toxicological Sciences</u> , 76:392-399 (2003).	<input type="checkbox"/>	
	221	KHAN et al., "Role of Serotonin in Fish Immunomodulation," <u>The Journal of Experimental Biology</u> , 200:1833-1838 (1997).	<input type="checkbox"/>	
	222	KHERSONSKY et al., "Facilitated Forward Chemical Genetics Using a Tagged Triazine Library and Zebrafish Embryo Screening," <u>J. Am. Chem. Soc.</u> , 125:11804-11085 (2003).	<input type="checkbox"/>	
	195	KURLANDSKY et al., "Plasma Delivery of Retinoic Acid to Tissues in the Rat," <u>J. Biol. Chem.</u> , 270(30):17850-17857 (1995).	<input type="checkbox"/>	
Examiner Signature				Date Considered

\* EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup> Applicant's unique citation designation number (optional). <sup>2</sup> Applicant is to place a check mark here if English language Translation is attached.

Substitute for form 1449B/PTO  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (Use as many sheets as necessary)		<b>Complete if Known</b>			
		Application Number	10/678,765		
		Filing Date	October 2, 2003		
		First Named Inventor	Serbedzija, George N.		
		Art Unit	1632		
		Examiner Name	Valarie E. Bertoglio		
Sheet	4	of	5	Attorney Docket Number	018852-000511US

NON PATENT LITERATURE DOCUMENTS			
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>
	193	LUNDQUIST, Frank. "The Blood-Brain-Barrier in Some Freshwater Teleosts," <u>Acta Physiologica Scandinavica</u> , 4(3-4):201-206 (1942).	<input type="checkbox"/>
	223	MACRAE et al., "Zebrafish-Based Small Molecule Discovery," <u>Chemistry &amp; Biology</u> , 10:901-908 (2003).	<input type="checkbox"/>
	199	MILAN et al., "Drugs that Induce Repolarization Abnormalities Cause Bradycardia in Zebrafish," <u>Circulation</u> , 107:1355-1358 (2003).	<input type="checkbox"/>
	192	MILLER et al., "Xenobiotic efflux pumps in insolated fish brain capillaries," <u>Am J Physiol Regulatory Integrative Comp Physiol</u> , 282:R191-198 (2002).	<input type="checkbox"/>
	207	MOORE et al., "Murine Cerebral Microvascular Endothelium Incorporate and Metabolize 12-Hydroxyeicosatetraenoic Acid," <u>Journal of Cellular Physiology</u> , 137:785-85 (1988).	<input type="checkbox"/>
	205	MOOS et al., "Cerebrovascular permeability to azo dyes and plasma proteins in rodents of different ages," <u>Neuropathology and Applied Neurobiology</u> , 19:120-127 (1993).	<input type="checkbox"/>
	229	Opposition Brief dated 01/11/2008 filed by opponent during opposition proceedings for EP 1 644 733.	<input type="checkbox"/>
	224	PEITSARO et al., "Modulation of the histaminergic system and behaviour by $\alpha$ -fluoromethylhistidine in zebrafish," <u>Journal of Neurochemistry</u> , 86:432-441 (2003).	<input type="checkbox"/>
	198	PRADEL et al., "Inhibition of Memory Consolidation by Antibodies against Cell Adhesion Molecules after Active Avoidance Conditioning in Zebrafish," <u>J. Neurobiol.</u> , 39:197-206 (1999).	<input type="checkbox"/>
	225	RENIER et al., "Genomic and functional conservation of sedative-hypnotic targets in the zebrafish," <u>Pharmacogenetics and Genomics</u> , 17:237-253 (2007).	<input type="checkbox"/>
	226	RITTER et al., "In Vivo Imaging of Zebrafish Reveals Differences in the Spinal Networks for Escape and Swimming Movements," <u>The Journal of Neuroscience</u> , 21(22):8956-8965 (2001).	<input type="checkbox"/>
Examiner Signature			Date Considered

\*EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>1</sup> Applicant's unique citation designation number (optional). <sup>2</sup> Applicant is to place a check mark here if English language Translation is attached.

Substitute for form 1449B/PTO  <b>INFORMATION DISCLOSURE STATEMENT BY APPLICANT</b>  (Use as many sheets as necessary)				<b>Complete if Known</b>	
				Application Number	10/678,765
				Filing Date	October 2, 2003
				First Named Inventor	Serbedzija, George N.
				Art Unit	1632
				Examiner Name	Valarie E. Bertoglio
Sheet	5	of	5	Attorney Docket Number	018852-000511US

NON PATENT LITERATURE DOCUMENTS				
Examiner Initials *	Cite No. <sup>1</sup>	Include name of the author (in CAPITAL LETTERS), title of the article (when appropriate), title of the item (book, magazine, journal, serial, symposium, catalog, etc.), date, page(s), volume-issue number(s), publisher, city and/or country where published.	T <sup>2</sup>	
	204	SKOV et al., "The blood volumes of the primary and secondary circulatory system in the Atlantic cod <i>Gadus morhua</i> L., using plasma bound Evans Blue and compartmental analysis," <u>The Journal of Experimental Biology</u> , 206:591-599 (2003).	<input type="checkbox"/>	
	203	STOPA et al., "Albumin binds self-assembling dyes as specific polymolecular ligands," <u>International Journal of Biological Macromolecules</u> , 40:1-8 (2006).	<input type="checkbox"/>	
	227	STUERMER et al., "Development of the Retinotectal Projection in Zebrafish Embryos under TTX-induced Neural -Impulse Blockade," <u>The Journal of Neuroscience</u> , 10(11):3615-3626 (1990).	<input type="checkbox"/>	
	228	SVOBODA et al., "Nicotinic Receptors Mediate Changes in Spinal Motoneuron Development and Axonal Pathfinding in Embryonic Zebrafish Exposed to Nicotine," <u>The Journal of Neuroscience</u> , 22(24):10731-10741 (2002).	<input type="checkbox"/>	
	201	VAN DE WATERBEEMD et al., "Estimation of Blood-Brain Barrier Crossing of Drugs Using Molecular Size and Shape, and H-Bonding Descriptors," <u>Journal of Drug Targeting</u> , 6(2):151-165 (1998).	<input type="checkbox"/>	
	196	ZHDANOVA et al., "Melatonin promotes sleep-like state in zebrafish," <u>Brain Research</u> , 903:263-268 (2001).	<input type="checkbox"/>	

Examiner Signature		Date Considered	
--------------------	--	-----------------	--

<sup>1</sup> EXAMINER: Initial if reference considered, whether or not citation is in conformance with MPEP 609. Draw line through citation if not in conformance and not considered. Include copy of this form with next communication to applicant.

<sup>2</sup> Applicant's unique citation designation number (optional). <sup>2</sup> Applicant is to place a check mark here if English language Translation is attached.